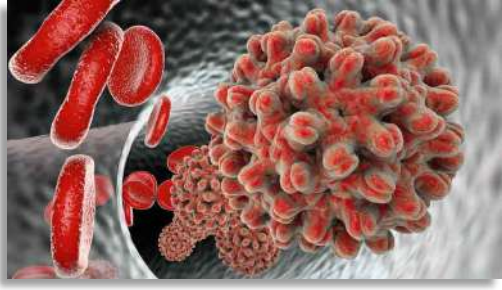


Klinik Bulgular ve Ekstrahepatik Tutulum

Dr.R.Aytaç ÇETİNKAYA



Akut Enfeksiyonu

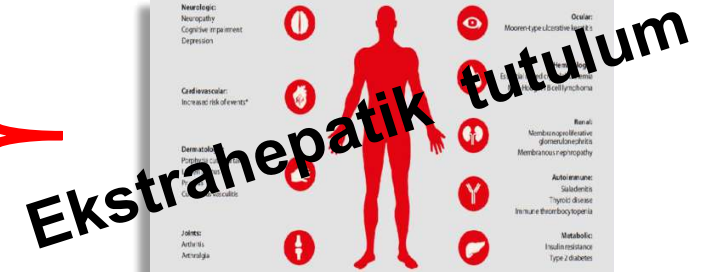
HCV RNA klirens (%15-25)

(Asemptomatik %70-80)

Fulminan (nadir)

Kronik Enfeksiyonu (%75-85)

Kronik Aktif Hepatit



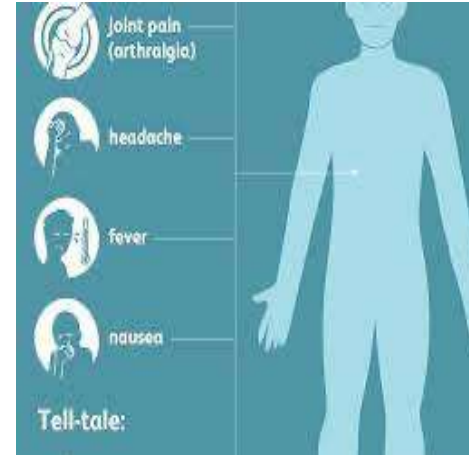
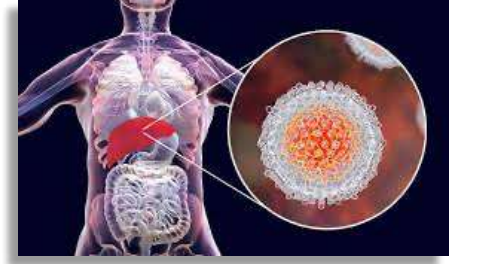
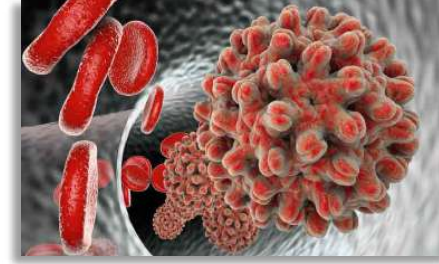
Dekompanse Siroz
5 yıl survi %50

Siroz (%10-20)
> 20 yıl

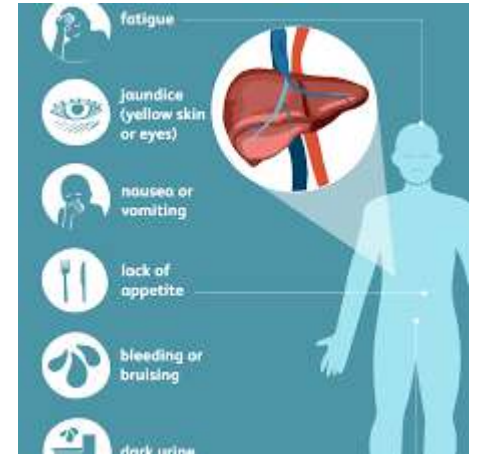
HCC %1-4 her yıl

Akut enfeksiyon klinik tablo

- Asemptomatik (%75)
- Semptomatik –bulgu (%25)
 - ✓ Eklem ağrısı
 - ✓ Baş ağrısı
 - ✓ Ateş
 - ✓ Mide bulantısı (sigara, yağlı yemek)
 - ✓ Karın ağrısı
 - ✓ Sarılık
 - ✓ Hepatoseplenomegali
 - ✓ Ciltte döküntü
 - ✓ Koyu renk idrar
 - ✓ Akolik gaita

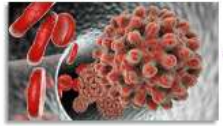


Tipik olmayan



Tipik bulgular

Kronik Enfeksiyon / Hepatit klinik tablo

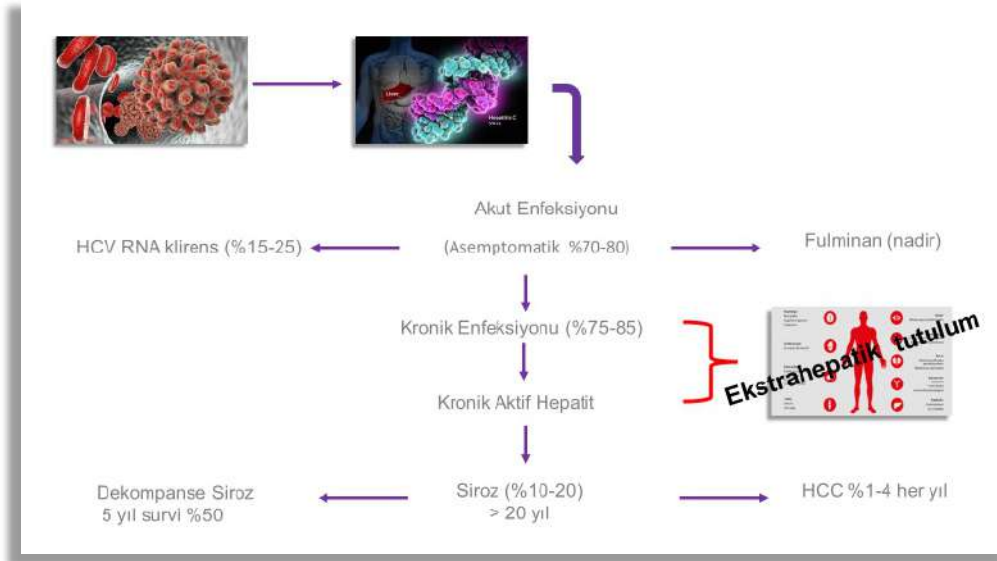


- Asemptomatik
- Semptomatik -bulgu

✓ Halsizlik

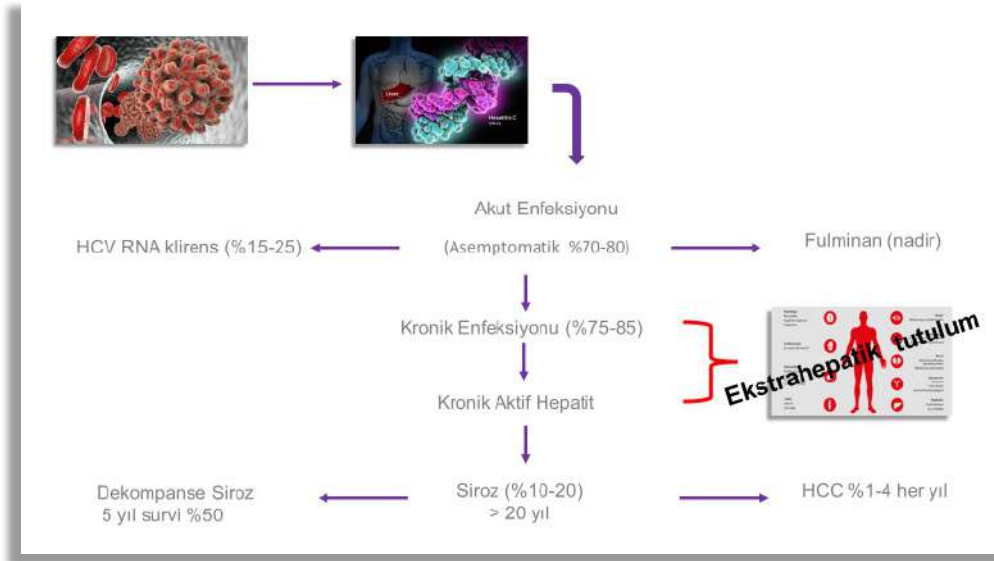
✓ Yorgunluk

Ekstrahepatik tutulum gözden kaçırıyor muyuz?



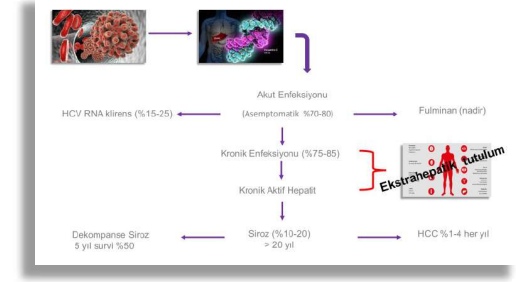
- The GERMIVIC
- Prospektif
- Çok merkezli
- 321 Kronik HCV enfeksiyon/hepatit
- %74
- En az bir ekstrahepatik bulgu

Ekstrahepatik tutulum kimlerde daha sık?



- Hasta yaşı ?
- **Hastalık yaşı**
- Kadın cinsiyet
- Karaciğer fibrozis

Ekstrahepatik hastalık en sık ?

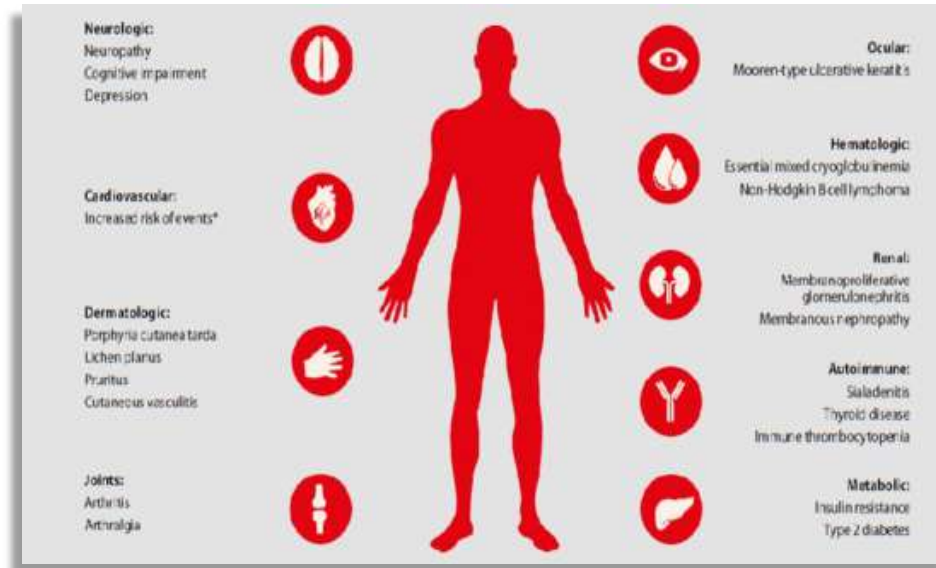
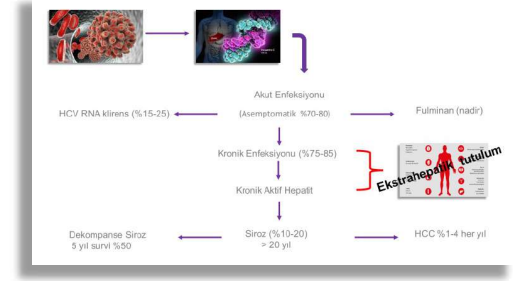


✓ Esansiyel mikst kriyoglobulinemi

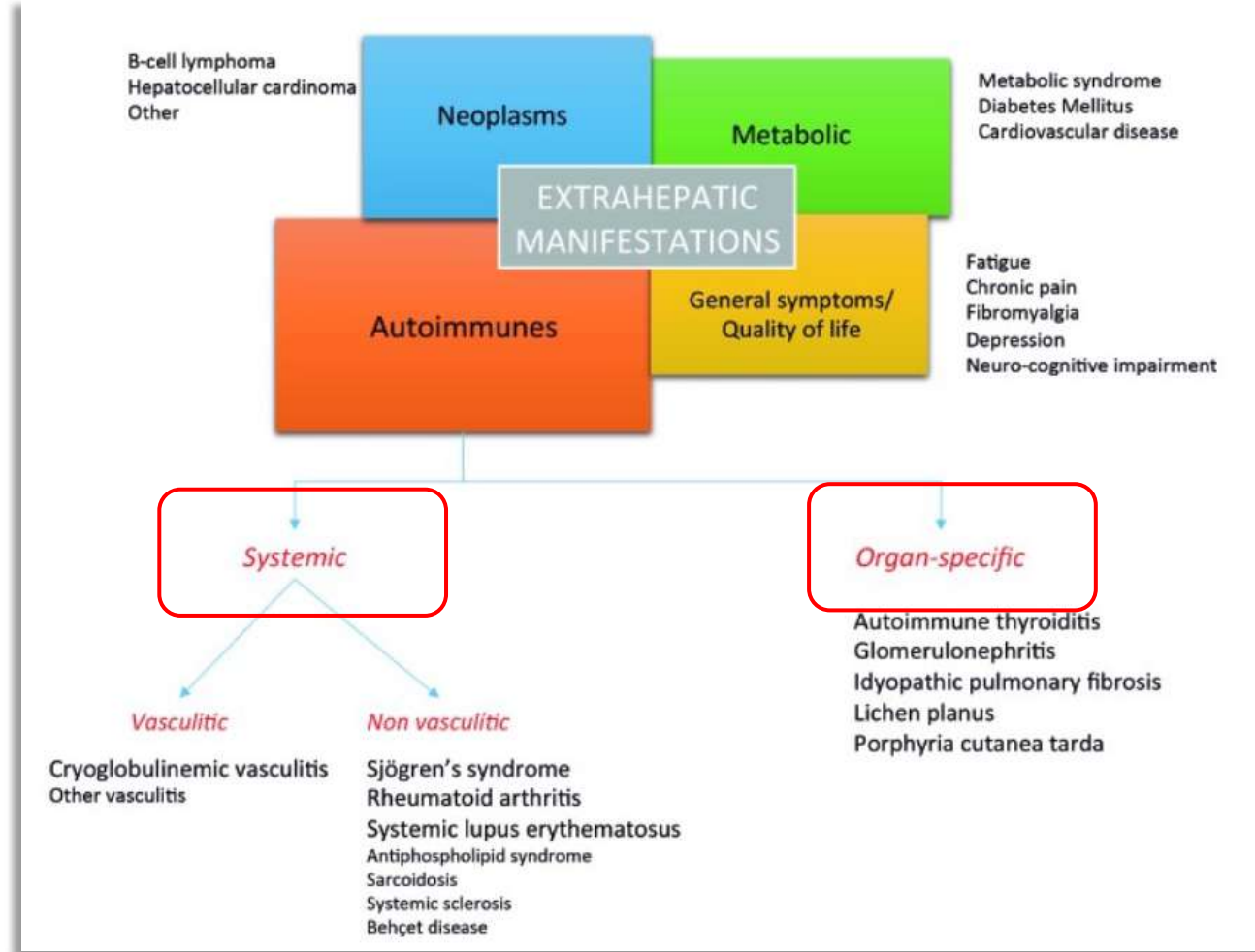
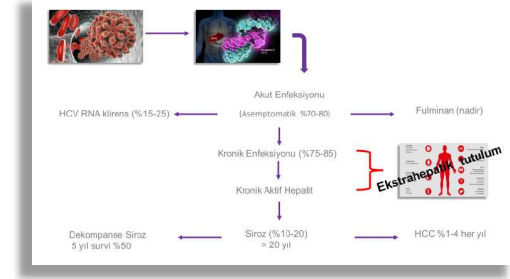
✓ Lenfoma



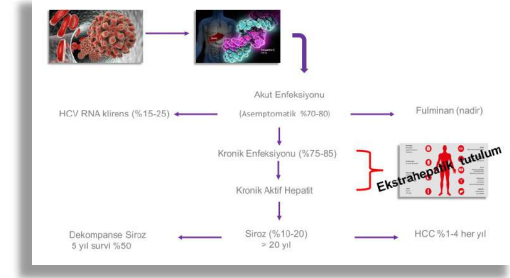
Ekstrahepatik hastalık



Ekstrahepatik hastalık



Ekstrahepatik hastalık



a) Mild and Moderate Manifestations

- Purpura
- Sporadic cutaneous ulceration
- Arthralgia/Arthritis
- Non-inflammatory muscle pain
- General conditions (malaise, fever)
- Mild to moderate neuropathy (sensitive)

b) Severe manifestations

- Persisting, multiple cutaneous ulceration that does not heal
- Finger ischemia
- Severe neuropathy (Motor or motor-sensitive)
- Glomerulonephritis with / without renal disease / nephrotic syndrome
- Interstitial lung disease
- vascular gastrointestinal affectation (non-necrotizing)
- severe autoimmune cytopenias (symptomatic hemolytic anemia / thrombocytopenia)

c) potentially fatal manifestations

- Rapidly progressive glomerulonephritis
- CNS affectation
- Acute Necrotizing intestinal vasculitis
- Alveolar hemorrhage
- Coronary artery disease (other etiology excluded)

Lenfoproliferatif hastalıklar

- Esansiyel miks kriyoglobulinemi
- Lenfoma
- Monoklonal gammopati

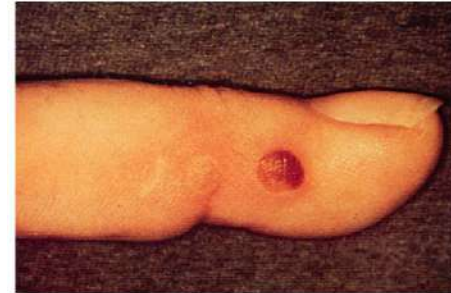
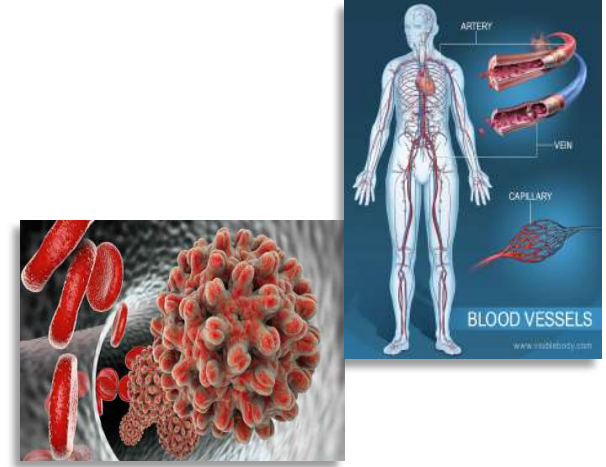
Esansiyel miks kriyoglobulinemi

- Tip II kriyoglobulinemi
- > %90 HCV
- küçük - orta büyük damar
- İmmunkompleks birikimi
- Membranoproliferatif **glomerülonefrit**
- Lökositoklastik vaskulit palpabl **purpura ve peteşi**

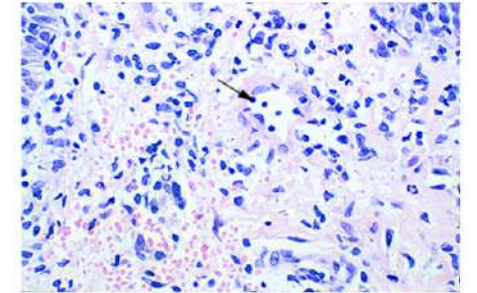


Esansiyel miks kriyoglobulinemi

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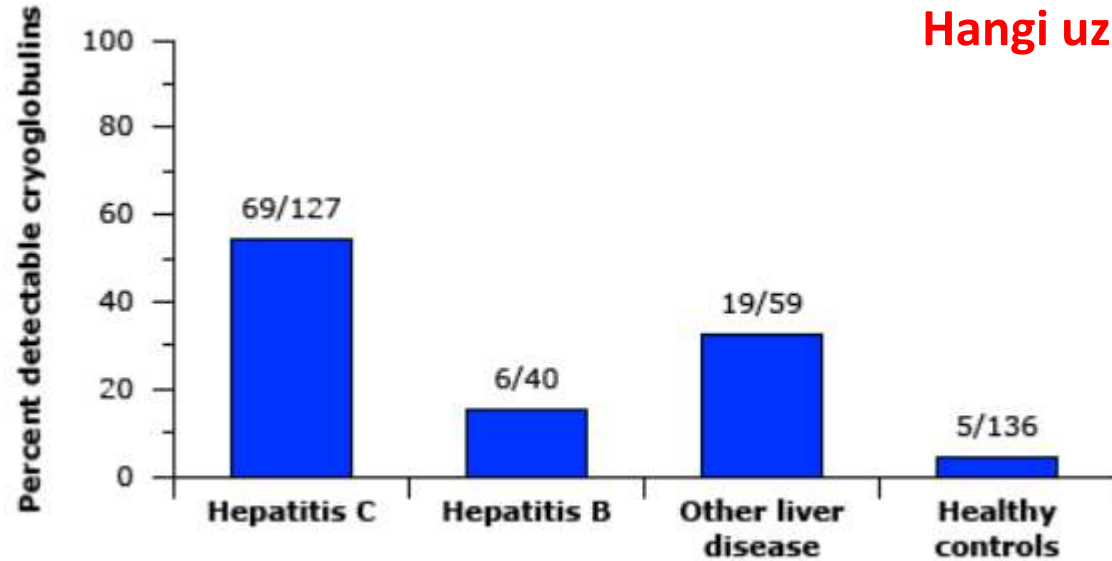


Leukocytoclastic vasculitis appearing as raised purpura. This lesion can occur with any vasculitic syndrome and in the collagen vascular diseases.

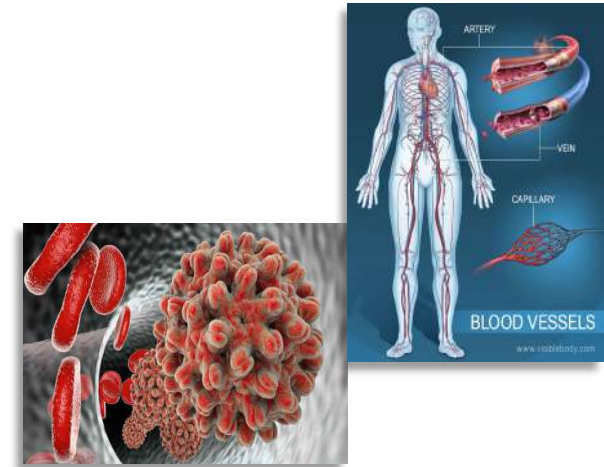





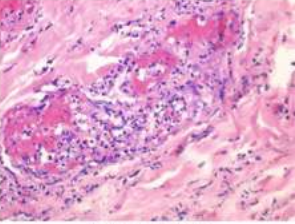
Leukocytoclastic vasculitis involving the dermal papillae capillaries and venules (arrow), a finding that probably reflects an Arthus type III immune complex reaction.

Esansiyel miks kriyoglobulinemi



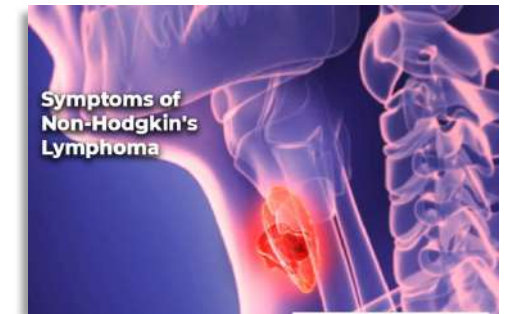
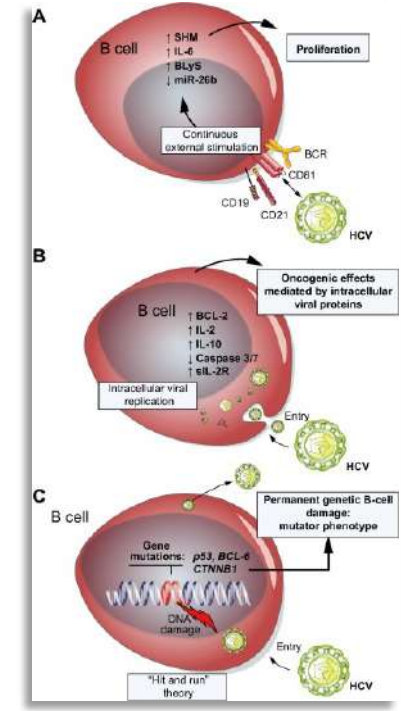
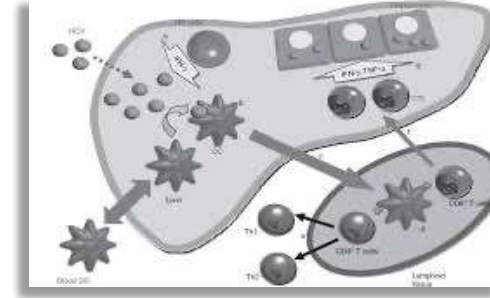
Tedavi ?
Hangi uzmanlık ?



Clinical	Serological	Pathological		
 <p>Recent purpura</p>	 <p>Dyschromic lesions</p>	 <p>Cryoprecipitate after 7 days, +4 °C</p>		
 <p>Leukocytoclastic vasculitis</p>				
<p>Serum MCs alone</p> <ul style="list-style-type: none"> • Possible preclinical condition with or without RF and/or low C4 • Careful clinical evaluation of possible underlying infectious (HCV or HBV) autoimmune and/or haematological and/or neoplastic disease • Monitoring without treatment 	<p>Mixed cryoglobulinaemia vasculitis</p> <table border="0"> <tr> <td> <p>Clinical</p> <ul style="list-style-type: none"> • Purpura • Weakness • Arthralgias • Liver involvement • Renal involvement • Skin involvement • Peripheral neuropathy </td> <td> <p>Serological</p> <ul style="list-style-type: none"> • Mixed cryoglobulins • RF+ • Low C4 <p>Pathological</p> <ul style="list-style-type: none"> • Leukocytoclastic vasculitis • B cell expansion </td> </tr> </table>	<p>Clinical</p> <ul style="list-style-type: none"> • Purpura • Weakness • Arthralgias • Liver involvement • Renal involvement • Skin involvement • Peripheral neuropathy 	<p>Serological</p> <ul style="list-style-type: none"> • Mixed cryoglobulins • RF+ • Low C4 <p>Pathological</p> <ul style="list-style-type: none"> • Leukocytoclastic vasculitis • B cell expansion 	<p>Clinical symptoms without serum MCs</p> <ul style="list-style-type: none"> • Absence of cryoprecipitable IC • Repeat cryoglobulin detection at different time intervals to confirm the diagnosis of cryoglobulinaemic vasculitis • Careful clinical evaluation of possible underlying infectious (for example, HCV, HBV) autoimmune and/or haematological and/or neoplastic disease
<p>Clinical</p> <ul style="list-style-type: none"> • Purpura • Weakness • Arthralgias • Liver involvement • Renal involvement • Skin involvement • Peripheral neuropathy 	<p>Serological</p> <ul style="list-style-type: none"> • Mixed cryoglobulins • RF+ • Low C4 <p>Pathological</p> <ul style="list-style-type: none"> • Leukocytoclastic vasculitis • B cell expansion 			

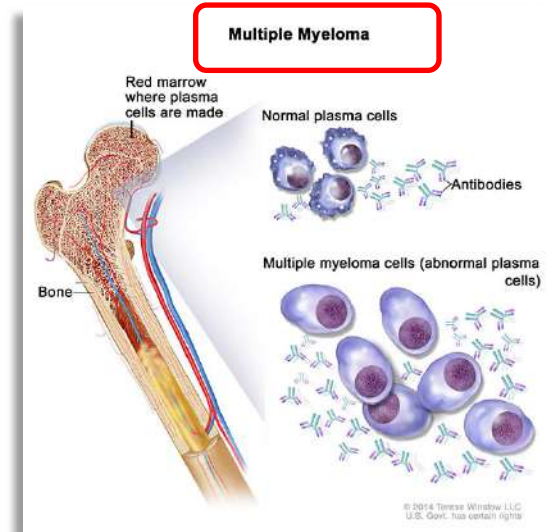
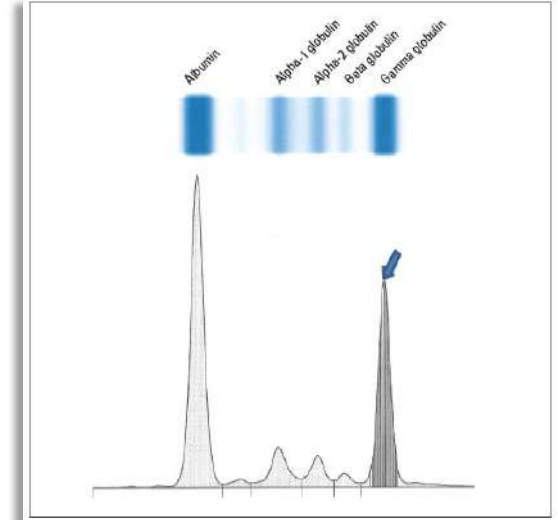
Lenfoma

- B hücreli non-hodgkin lenfoma
- 48 çalışma metaanaliz, 5542 hasta
 - > %15 HCV
 - Genel popülasyon %1,5
- Geniş kapsamlı 146,394 kohort çalışma
 - 7 yıl
 - 1,28 risk
 - 28 kat artış



Monoklonal gammopatiler

- HCV - B lenfosit – plazma hücresi
- 2 merkez vaka-kontrol
 - 239 HCV enfeksiyon
 - 98 HCV negatif (kontrol)
 - 26 monoklonal gammapati
 - Multiple myelom n:9

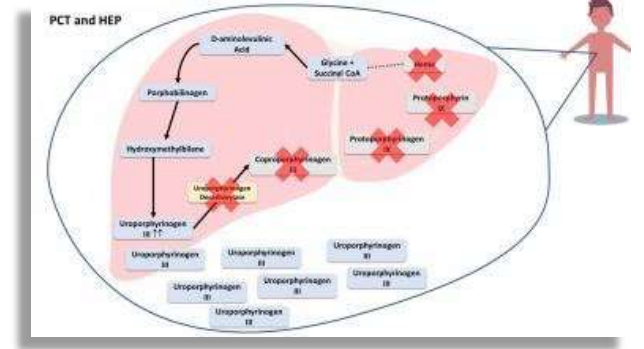


Dermatolojik hastalıklar

- Porfiriya kutanea tarda (PCT)
- Liken Planus
- Nekrolitik akral eritem
-
-

Dermatolojik Hastalıklar; Porfiriya kutanea tarda (PCT)

- Hepatik üroporfirinojen dekarboksilazın (UROD)
- Karaciğerde, kan ve idrarda **üroporfirinlerin birikmesi**
- **Veziküler**
- **Ciltte eroze lezyon**, ışığa duyarlı
- Güneşe maruz kalma ve/veya minör travma hemorajik
- 2167 PCT hasta 50 çalışmanın sistematik incelemesi
- HCV prevalansı yüzde 50
- **Antiviral tedavi; HCV viremi kaybolması PCT korele**



Dermatolojik Hastalıklar; Liken Planus

- Düz tepeli, morumsu, kaşıntılı papüller
- Mukoza mebran, Tırnak
- Sistematik inceleme,
 - ✓ Oral liken planus hastaları-kontrol grup
 - ✓ Reaktif anti-HCV x2, x6
- Japonya 261 Kr. HCV enfeksiyonu
 - ✓ 71 oral mukoza liken planus
 - ✓ x10



Otoimmün Bozukluklar

- Sublinik (ANA, AMA, RF, Antitiroglobulin, ASMA, Antikardiyolipin, LKM..)
- Tiroid hastalıklar
- Sialadenit
- Otoimmün trombositopenik purpura
- Sjögren sendromu (%48)
- SLE (%13)
- Poliartris Nodosa (%6)
- ..
-

Sjögren sendromu/Sicca semptomları

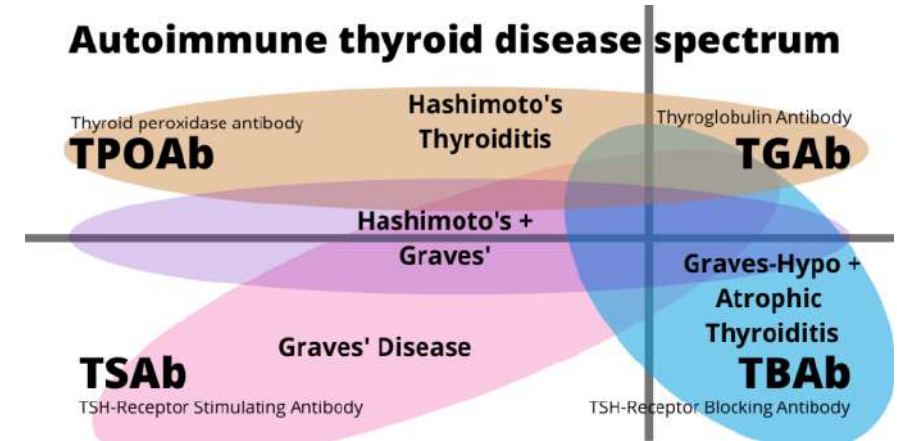
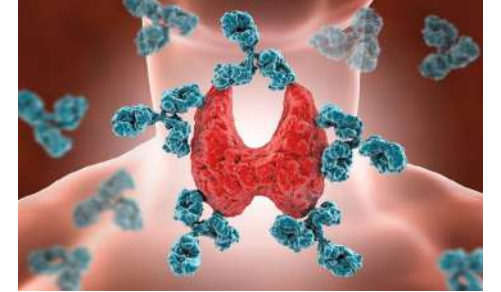
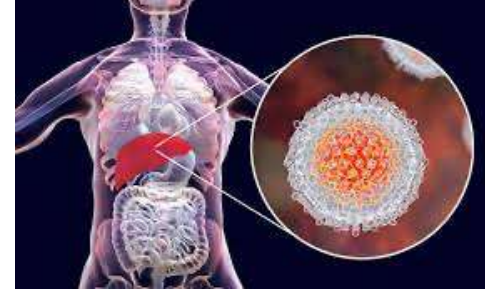
- Göz kuruluğu
- Dil kuruluğu
- Artrit
- Kuru cilt-Döküntü
- Sindirim problemleri
- Kuru öksürük
- 102 çalışma içeren metaanaliz
 - ✓ HCV – Sjögren %15
 - ✓ Kontrol <%1

Sjogren's Syndrome

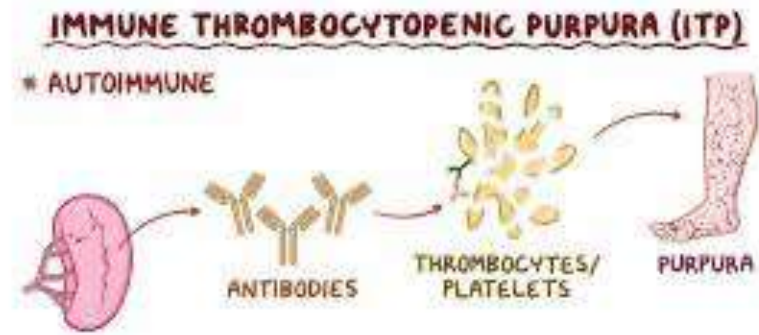


Tiroid hastalıkları

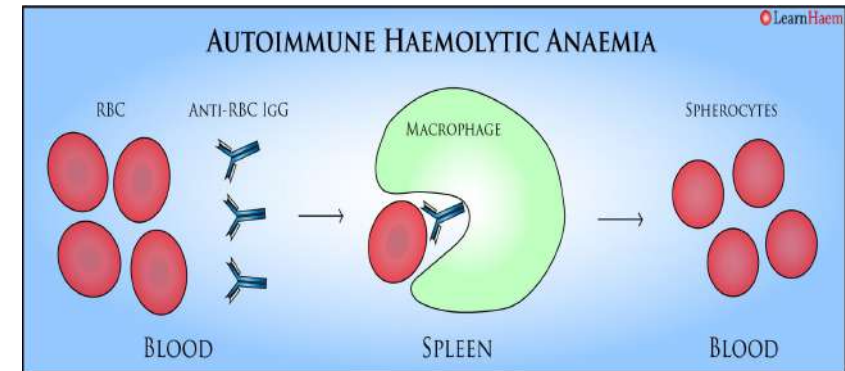
- HCV enfeksiyon
- Anti-tiroid antikor %5-17
- Hashimoto
- Graves hastalığı
- Tiroid hastalığı %2-13



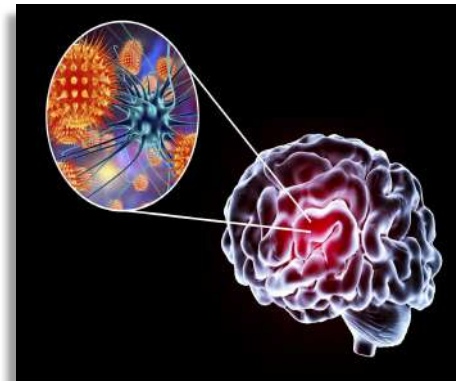
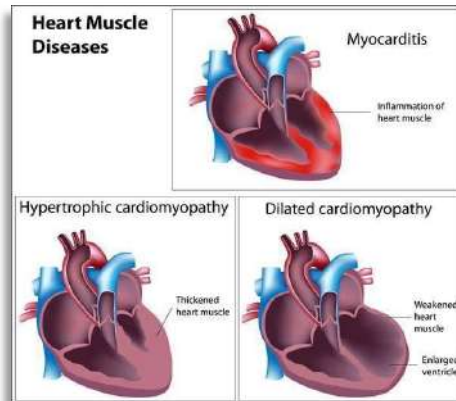
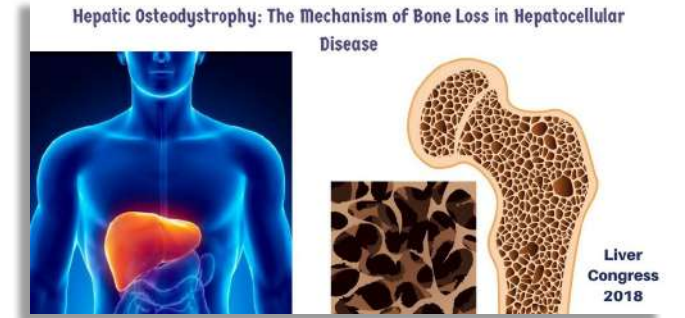
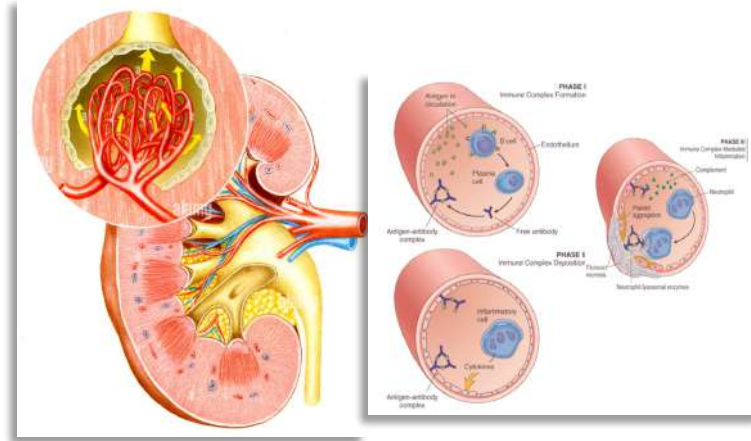
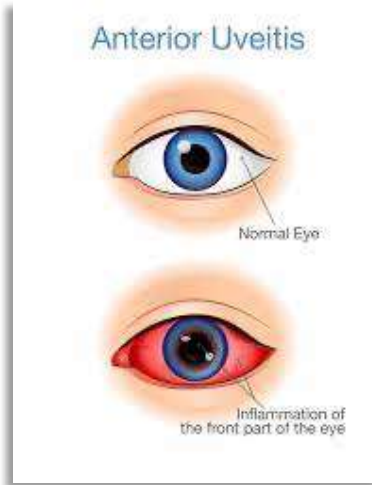
İmmun Trombositopeni(I TP),

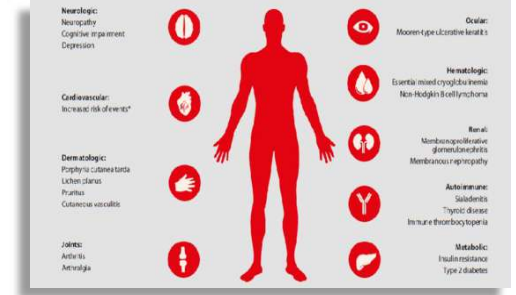
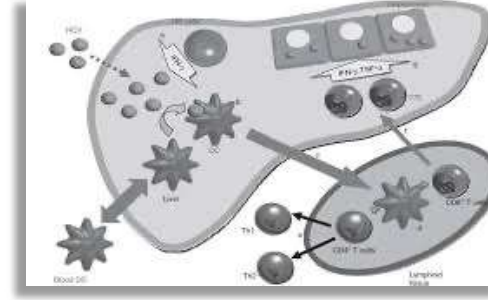
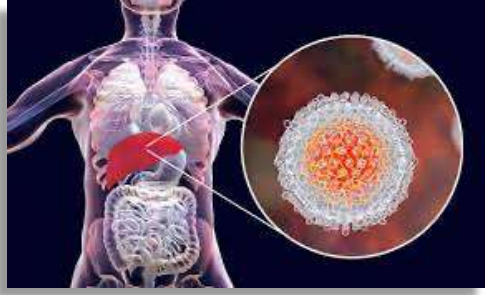


Otoimmun Hemolitik Anemi



Diğer tutulumlar





Klinik Bulgular ve Ekstrahepatik Tutulum

Dr.R.Aytaç ÇETİNKAYA